Supplementary Material 2. Certainty of Evidence Assessment Using GRADE Approach

Outcome	No. of Studies	Risk of Bias	Inconsistency	Indirectness	Imprecision	Publication Bias	Overall Certainty of Evidence
Sensitivity of combined biomarker–clinical models	17	no serious risk of bias	no serious inconsistency	not serious	not serious	Detected	⊕⊕⊕ Moderate
Specificity of combined biomarker–clinical models	16	no serious risk of bias	no serious inconsistency	not serious	not serious	Detected	⊕⊕⊕ Moderate
Diagnostic accuracy compared with single biomarkers	11	no serious risk of bias	very serious inconsistency	not serious	very serious	Detected	⊕ Very Low

⊕⊕⊕⊕ = High certainty of evidence; ⊕⊕⊕◯ = Moderate certainty of evidence; ⊕⊕◯◯ = Low certainty of evidence; ⊕◯◯◯ = Very low certainty of evidence